Liver Disease in the HIVinfected Patient

David Thomas, MD

Liver Disease and HIV Infection

- Spectrum of liver disease
- Hepatitis C
- Hepatitis B

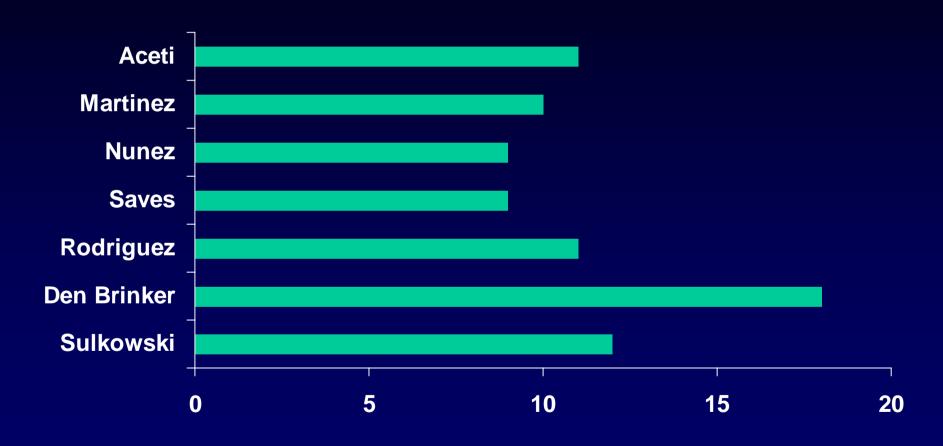
Opportunistic infections

- Opportunistic infections
 - Non TB mycobacteria
 - Lymphoma

- Opportunistic infections
- HIV related cholangitis
 - Cryptosporidium, MAI, CMV, microsporidia, HIV

- Opportunistic infections
- HIV related cholangitis
- Antiretroviral toxicity

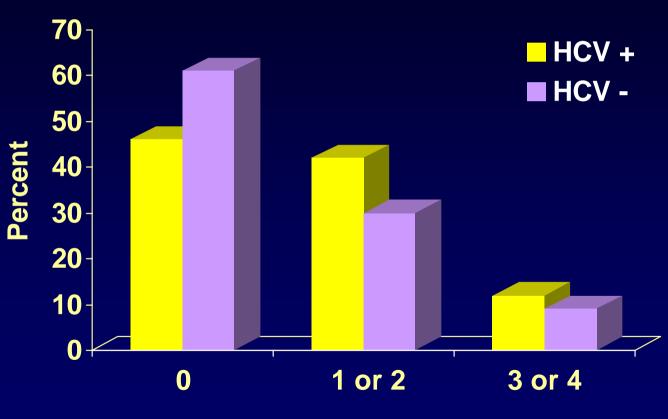
Grade 3-4 ALT/AST Elevations Following New HAART



Adapted from Soriano et al AIDS 2002:813

Incidence (%)

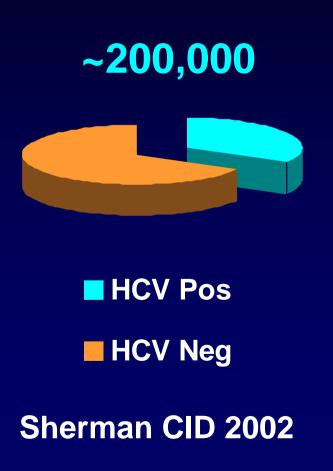
Hepatic Toxicity Grade by HCV Status Johns Hopkins HIV Cohort

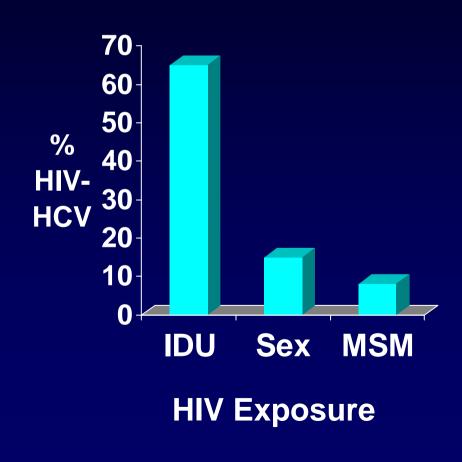


Hepatotoxicity Grade

Sulkowski JAMA 2000;283:2526

Hepatitis C in HIV-infected Persons in the USA



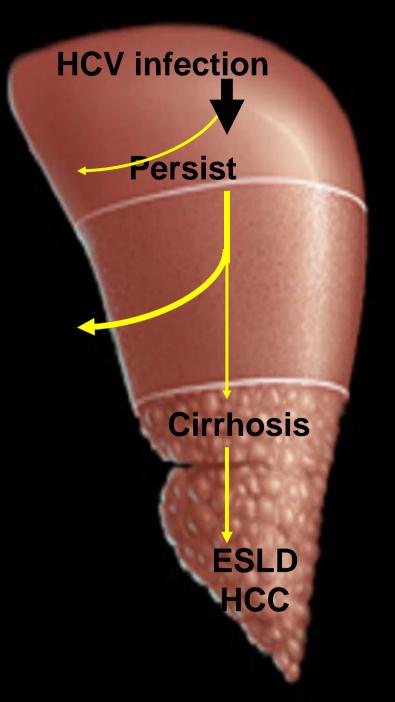


Hepatitis C Natural History

Effect of HIV

Recover

Silent infection

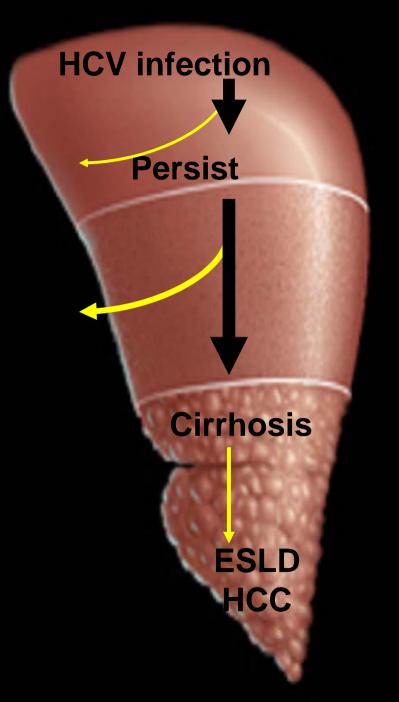


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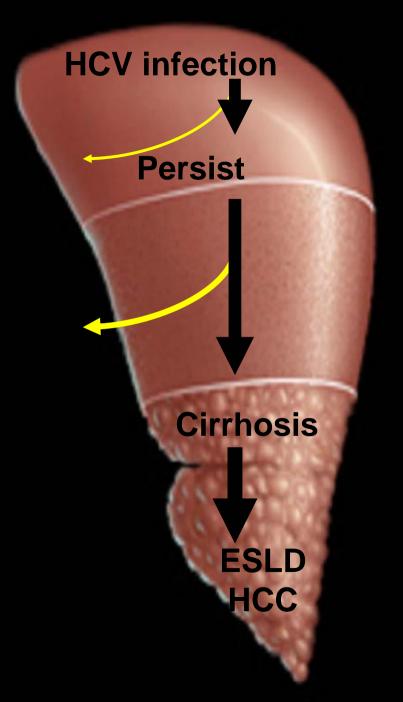


Hepatitis C Natural History

Effect of HIV

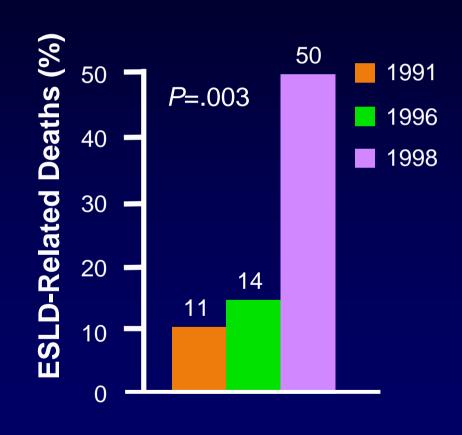
Recover

Silent infection

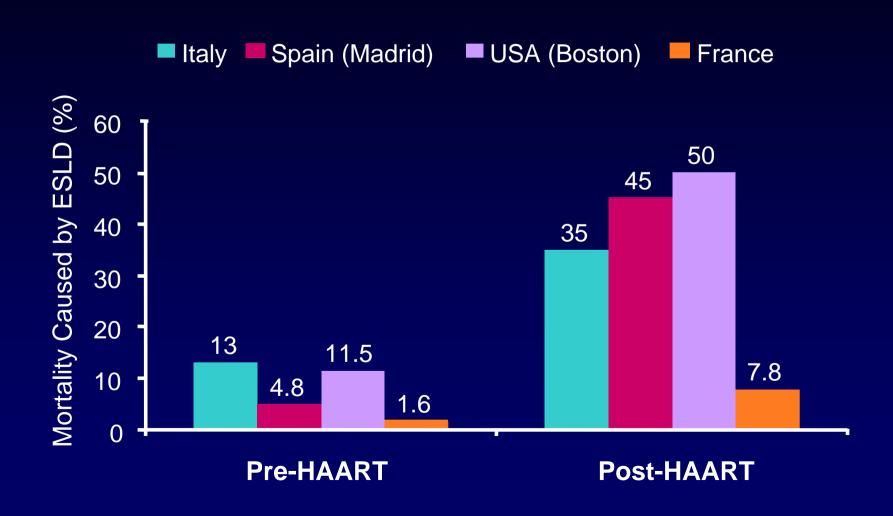


Increasing Mortality From ESLD in Patients With HIV

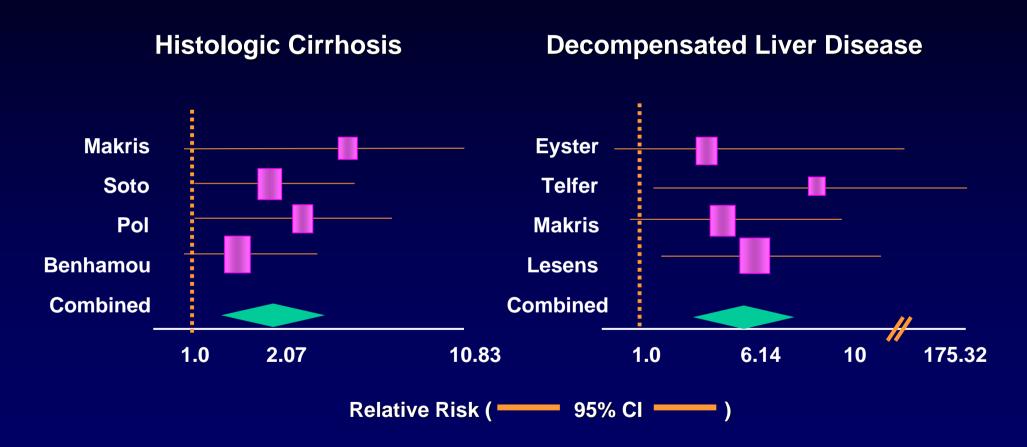
- 1/3 of 1998 cohort: discontinuing HAART due to hepatotoxicity
- >1/2 who died with ESLD: HIV RNA undetectable or CD4 >200/mm³



HAART and End Stage Liver Disease (ESLD) Mortality in HIV-infected Persons



Increased Risk of Cirrhosis and ESLD in HIV-HCV Coinfected Patients

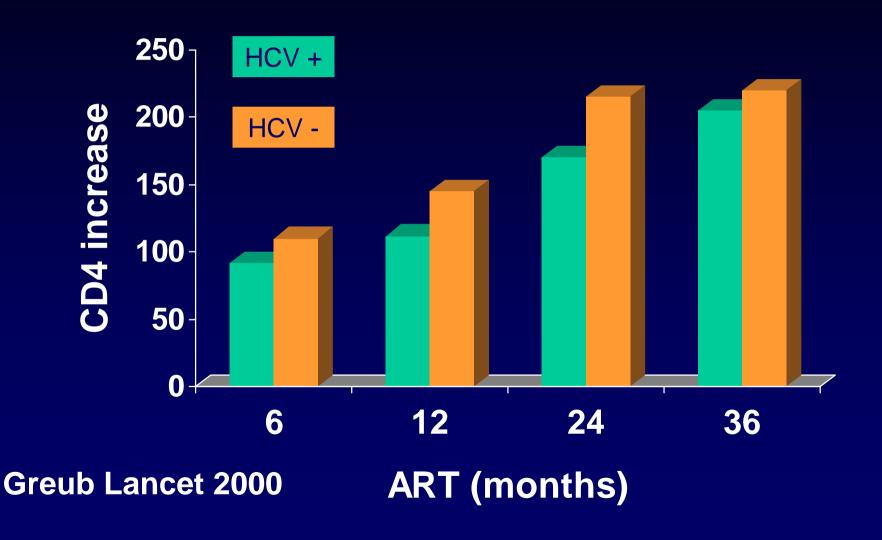


Graham C et al. CID 2001, Makris Br J Haematol 1996, Soto J Hepatol 1997, Pol J Hepatol 1998, Benhamou Hepatology 1999, Eyster JAIDS 1993, Telfer Br J Haematol 1994, Lesens JID 1999.

Outcome of HIV Infection According to HCV: Swiss HIV Cohort Study

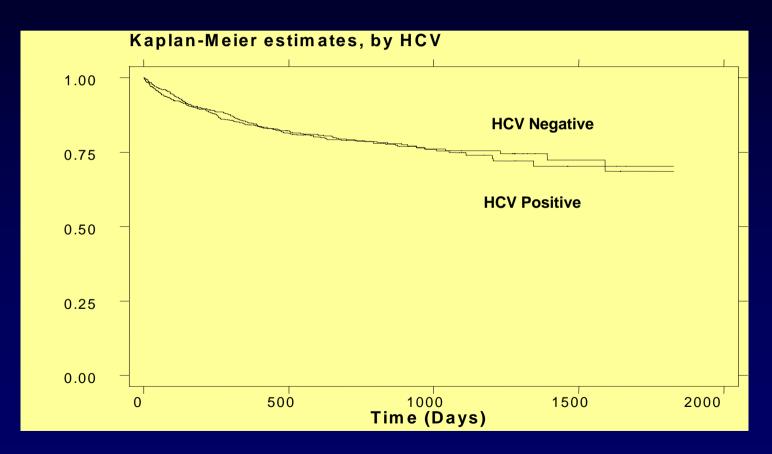
- 3111 of 3655, started HAART 6/96 5/99
- 1157 (37%) HCV positive
- Progression to death or AIDS: Haz = 1.7
- HIV VL <400: 1068 HCV neg and 528 HCV pos, Haz = 3.5
- CD4+ lymphocyte response

Suppressed CD4+ Lymphocyte Responses in HCV-HIV Coinfected Persons



Probability of Survival Without an Opportunistic Infection

1,742 pts at risk \rightarrow 328 events



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 - Prevalence of chronic hepatitis B
 8-10% in MACS and EuroSida Cohorts

Increased Liver Mortality in HIV-HBV Coinfected Men

- 5293 men (326 HBsAg+ baseline) followed to 3/00
- Overall liver MR 1.1/1000 PYs
- RR of liver death 18.7 in coinfected vs. only HBsAg+

16-14.2 14-Liver MR/1000 PYs **12**-10-8 6 8.0 1.7 2-0.0

Thio et al, The Lancet, 2002

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Important Areas for Future Research

- Pathogenesis
- Fibrosis progression rate
- Treatment outcomes and duration
- Liver transplantation
- Treatment access